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hm the
THERNATIONAL PRELIMINARY EXAMINING AUTHORITY

To:

KREMER, Simon, M.

Mewburn Ellis

York House
23 Kingsway

London WC2B 6HP

GRANDE BRETAGNE

RECEIVED

WRITTEN OPINION (PCT Rule 66)

15 AUG 2003 MEWBURN ELLIS

Date of malling (day/month/year)

13.08.2003

Applicant's or agent's file reference SMK/LP6138390

REPLY DUE

within 3 month(s) from the above date of mailing

International application No. PCT/EP02/14512

International filing date (day/month/year) 18.12.2002

Priority date (day/month/year)

19.12.2001

International Patent Classification (IPC) or both national classification and IPC

C12N15/82

Applicant

DŪRING, Klaus, et al.

- 1. This written opinion is the first drawn up by this International Preliminary Examining Authority.
- 2. This opinion contains indications relating to the following items:
  - I Basis of the opinion
  - II Priority
  - III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability
  - IV D Lack of unity of invention
  - V A Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
  - VI 

    Certain documents cited
  - VII 

    Certain defects in the international application
  - VIII 

    Certain observations on the international application
- 3. The applicant is hereby invited to reply to this opinion.

When? See the time limit indicated above. The applicant may, before the expiration of that time limit,

request this Authority to grant an extension, see Rule 66.2(d).

How? By submitting a written reply, accompanied, where appropriate, by amendments, according to Rule 66.3.

For the form and the language of the amendments, see Rules 66.8 and 66.9.

Also: For an additional opportunity to submit amendments, see Rule 66.4.

For the examiner's obligation to consider amendments and/or arguments, see Rule 66.4 bis.

For an Informal communication with the examiner, see Rule 66.6.

If no reply is filed, the international preliminary examination report will be established on the basis of this opinion.

 The final date by which the international preliminary examination report must be established according to Rule 69.2 ls: 19.04.2004

Name and mailing address of the international preliminary examining authority:



European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d Fax: +49 89 2399 - 4465 Authorized Officer

Grosskopf, R

Formalities officer (incl. extension of time limits)

Faux, K

Telephone No. +49 89 2399-8062



## WRITTEN OPINION

International application No.

PCT/EP02/14512

| -  |       |        |          |
|----|-------|--------|----------|
| I. | Pacie | AT THA | opinion  |
|    | حاجهط |        | Ophilion |

| ١. | With regard to the elements of the international application (Replacement sheets which have be        | en furnished to      |
|----|---|----------------------|
|    | the receiving Office in response to an invitation under Article 14 are referred to in this opinion as | "originally filed"): |

|   | Des   | Description, Pages                                     |   |  |  |  |  |  |
|---|---|--|---|--|--|--|--|--|
|   | 1-1   | 9  | as originally filed   |  |  |  |  |  |
|   | Cla   | ims, Numbers   |   |  |  |  |  |  |
|   |   |  |   |  |  |  |  |  |
|   | 1-1   | 2  | as originally filed   |  |  |  |  |  |
|   | Dra   | wings, Figures   |   |  |  |  |  |  |
|   | 1-2   |  | as originally filed   |  |  |  |  |  |
|   |   |  |   |  |  |  |  |  |
| 2.  | Witl<br>lang  | h regard to the langu<br>guage in which the in         | lage, all the elements marked above were available or furnished to this Authority in the ternational application was filed, unless otherwise indicated under this item. |  |  |  |  |  |
|   | The   | ese elements were av                                   | railable or furnished to this Authority in the following language: , which is:  |  |  |  |  |  |
|   |   | the language of a tra                                  | anslation furnished for the purposes of the international search (under Rule 23.1(b)).  |  |  |  |  |  |
|   |   | the language of pub                                    | lication of the international application (under Rule 48.3(b)).   |  |  |  |  |  |
|   |   | the language of a tra<br>Rule 55.2 and/or 55.          | anslation furnished for the purposes of international preliminary examination (under 3).  |  |  |  |  |  |
| 3.  | With<br>inte  | h regard to any <b>nucl</b> e<br>rnational preliminary | eotide and/or amino acid sequence disclosed in the international application, the examination was carried out on the basis of the sequence listing:                     |  |  |  |  |  |
|   | ☐ contained in the international application in written form. |  |   |  |  |  |  |  |
| filed together with the intern  |   |  | e international application in computer readable form.  |  |  |  |  |  |
|   |   | furnished subsequer                                    | ntly to this Authority in written form.   |  |  |  |  |  |
|   |   | furnished subsequer                                    | ntly to this Authority in computer readable form.   |  |  |  |  |  |
|   |   | The statement that t in the international a            | the subsequently furnished written sequence listing does not go beyond the disclosure application as filed has been furnished.  |  |  |  |  |  |
| The statement that the information recorded in computer readable form is identical to the written s listing has been furnished. |   |  |   |  |  |  |  |  |
| 4. The amendments have resulted in the cancellation of:   |   |  |   |  |  |  |  |  |
|   |   | the description,                                       | pages:  |  |  |  |  |  |
|   |   | the claims,  | Nos.:   |  |  |  |  |  |
|   |   | the drawings,  | sheets:   |  |  |  |  |  |
|   |   |  |   |  |  |  |  |  |

## **WRITTEN OPINION**

International application No.

PCT/EP02/14512

| 5. 🗆 | This opinion h | ıas been e  | established | as if (some | e of) the | amendments      | had not been | made, si | ince they | have |
|------|----------------|-------------|-------------|-------------|-----------|-----------------|--------------|----------|-----------|------|
|      | been consider  | red to go b | peyond the  | disclosure  | as filed  | (Rule 70.2(c)). | •            |          | -         |      |

(Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this opinion.)

- 6. Additional observations, if necessary:
- V. Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- 1. Statement

. )

Novelty (N)

Claims

Inventive step (IS)

Claims

1-12

Industrial applicability (IA)

Claims

2. Citations and explanations

see separate sheet

6-11-04; 1:3/AM

( )

## Ad item V:

The present claims are directed to a method for increasing the content of a transgene-encoded biomolecule in an organism, characterised by changing the distribution of ATP and/or ADP in cells of the organism. The expression "changing the distribution of ATP and/or ADP" is merely the characterisation of the essential feature by the desired result to be achieved without indicating the means how said change should be carried out. In addition, said expression neither indicates the extent of the change nor the exact localisation where said change should take place ("in cells").

In view of this vague and broad definition of the main feature and when taking into account that nearly each reaction in the cell involves a "change of distribution of ATP" (e.g. the use of glucose in a culture medium), as indicated in the search report, a meaningful search for an accordingly characterised claim over the whole range was not possible.

As a consequence, also the examination has to be limited to the those means which were used in the present application in order to achieve the change of ATP/ADP distribution, i.e. the use of the cloned plastidiary ATP/ADP transporter.

The quoted documents are:

- (1) WO 99 58654 A (MAX PLANCK GESELLSCHAFT; MOEHLMANN TORSTEN (DE); MARTINI NORBERT () 18 November 1999 (1999-11-18)
- (2) GEIGENBERGER P ET AL: "OVEREXPRESSION OF PYROPHOSPHATASE LEADS TO INCREASED SUCROSE DEGRADATION AND STARCH SYNTHESIS, INCREASED ACTIVITIES OF ENZYMES FOR SUCROSE-STARCH INTERCONVERSIONS, AND INCREASED LEVELS OF NUCLEOTIDES IN GROWING POTATO TUBERS" PLANTA, SPRINGER VERLAG, DE, vol. 205, no. 3, July 1998 (1998-07), pages 428-437, XP000997825 ISSN: 0032-0935

The system which has been used in the present application to change the distribution of ATP has been described in D1.

Moreover, from several of the documents cited in the search report it is known that a change of ATP has an influence on various activities, including the increase of enzyme activities and the increase of the content of various compounds (see D2). Thus, a skilled person would certainly not be surprised that a "change" in ATP

WRITTEN OPINION SEPARATE SHEET

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International application No. PCT/EP02/14512

concentration, in general, may also have an influence e.g. on the expression of a transgene which may result in a higher or lower content of said transgene.

Therefore, the general disclosure of the claims, even when taking into account of the specific system which has been used to change the ATP distribution in a cell (see above), cannot be regarded as involving an inventive activity.